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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

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Patentee: Serial No: Milanovski, et al. 09/763,345

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August 24, 1999

For:

METHOD OF ELECTROCHEMICAL ANALYSIS OF AN ANALYTE

CERTIFICATE OF MAILING UNDER 37 C.F.R. §1.8(a)

The undersigned hereby certifies that this document is being placed in the United States mail with first-class postage attached, addressed to Commissioner for Patents, P.O. Box_1450, Alexandria, VA 22313-1450, on the GTZ, day of October, 2004.

Mail Stop Certificate of Correction

Commissioner for Patents

P.O. Box 1450

Alexandria, VA 22313-1450

REQUEST FOR ENTRANCE OF CERTIFICATE OF CORRECTION UNDER 35 U.S.C. §254 and §255

Sir/Madam:

Patentee respectfully requests the correction of errors in the printing of the above-captioned patent. Specifically, claim 40 has typographical errors made by the Patent Office. Please correct as follows: In column 40, line 9, the second occurrence of "the" should be deleted.

Patentee points out that the corrections requested does not involve change in the patent that constitutes new matter or would require reexamination, and therefore, respectfully request that a certificate of correction be issued. Patentee encloses a copy of the issued patent with the errors highlighted. Since the error was made by the Patent Office, it is respectfully submitted that no fee is due. However, if the Examiner deems a fee necessary, the fee may be charged to the account of the undersigned, Deposit Account No. 23/2825. Should any questions arise concerning the foregoing, please contact the undersigned at the telephone number listed below.

For the reasons stated above, Patentee respectfully requests entrance of the enclosed Certificate of Correction.

> Respectfully submitted, Milanovski, et al., Patentee

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Docket No. B0192.70026US00 Date: October 6, 2004

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UNITED STATES PATENT AND TRADEMARK OFFICE

CERTIFICATE OF CORRECTION

PATENT NO.:

US 6,770,190 B1

DATED

August 3, 2004

INVENTORS:

Milanovski, et al.

It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

should read:

In Column 40, line 9, delete the second occurrence of "the".

Approved. 03/01/05

MAILING ADDRESS OF SENDER:

PATENT NO. US 6,770,190 B1

John R. Van Amsterdam Wolf, Greenfield & Sacks, P.C. 600 Atlantic Avenue Boston, Massachusetts 02210-2211

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- (a) providing a sensing electrode having an electroconductive polymer coating, the coating having immobilized therein or adsorbed thereto receptors which are capable of binding to the desired analyte to be detected in the sample;
- (b) contacting the sensing electrode with a test solution comprising the sample so that the said analyte binds to said immobilized or adsorbed receptors;
- (c) contacting the sensing electrode with a solution comprising secondary receptors capable of binding to said analyte at a site spatially distinct from the site of binding to immobilized or adsorbed receptors, said secondary receptors being conjugated with an enzyme;
- (d) monitoring the electric potential difference between the sensing electrode of part (c) and a reference electrode when both are immersed in an electrolyte; and
- (e) monitoring the electric potential difference between the sensing electrode of part (d) and a reference electrode following exposure to an electrolyte comprising 20 the substrate for said enzyme.

34. A method as claimed in claim 33 wherein the enzyme is capable of converting a substrate which has no detectable effect on the redox composition of the electroconductive polymer coating of the sensing electrode to a product capable of directly or indirectly affecting the redox composition of the said electroconductive polymer coating.

35. A method as claimed in claim 34 wherein the enzyme is a peroxidase.

36. A method as claimed in claim 34 wherein the product 30 capable of indirectly affecting the redox composition of the electroconductive polymer membrane causes a change in the pH of the electrolyte of part (e).

37. A method as claimed in claim 36 wherein the enzyme

38. A method as claimed in claim 33 wherein the enzyme is capable of converting a substrate which has no detectable effect on the redox composition of the electroconductive polymer coating of the sensing electrode to a product which is a substrate for a second enzyme, the action of the second enzyme generating a second product which directly or indirectly affects the redox composition of the electroconductive polymer coating of the sensing electrode.

39. A method as claimed in claim 33 wherein the enzyme is capable of converting a substrate which directly affects the 45 redox composition of the electroconductive polymer coating of the sensing electrode to a product which has no detectable effect on the redox composition of the said electroconductive polymer coating.

40. A method of electrochemical detection of an analyte in a sample, which method comprises the steps of:

- (a) providing a sensing electrode having an electroconductive polymer coating, the coating having immobilized therein or adsorbed thereto receptors which are capable of binding to the desired analyte to be detected in the sample;
- (b) contacting the sensing electrode with a test solution comprising the sample so that the the desired analyte to be detected in the sample binds to said immobilized or adsorbed receptors;
- (c) contacting the sensing electrode with a solution comprising competing molecules capable of binding to said immobilized or adsorbed receptors, said competing molecules being conjugated with an enzyme;
- (d) monitoring the electric potential difference between the sensing electrode of part (c) and a reference electrode when both are immersed in an electrolyte; and
- (e) monitoring the electric potential difference between the sensing electrode of part (d) and a reference electrode following exposure to an electrolyte comprising the substrate for said enzyme.

41. A method of electrochemical detection of an analyte in a sample, which method comprises the steps of:

- (a) providing a sensing electrode comprising an electrically conductive electrode coated with a layer of electroconductive polymer with molecules of avidin or streptavidin immobilized therein or adsorbed thereto, said avidin or streptavidin molecules being attached to receptor molecules capable of binding the analyte to be detected attached via a biotin/avidin or biotin/streptavidin binding interaction;
- (b) contacting the sensing electrode with a test solution comprising the sample so that said desired analyte binds to said immobilized or adsorbed receptor molecules;
- (c) monitoring the potential of the sensing electrode relative to a reference electrode when both are immersed in an electrolyte; and
- (d) monitoring the potential difference of the sensing electrode relative to the reference electrode following a change in the ionic strength or composition of the electrolyte at constant pH.
- 42. A method as claimed in claim 41 wherein the analyte to be detected is a nucleic acid and the receptor molecules are oligonucleotides.